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14 April 2005

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PCT SPECIAL

Attorney Docket No. P51375

PROGRAMS OFFICE

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant:

Behrens, et al.

14 April 2005

Int'l Serial No.: PCT/US03/28654

Art Unit: unknown

Int'l Filing Date: 12 September 2003

Examiner: unknown

For:

A SET OF UBIQUITOUS CELLULAR PROTEINS INVOLVED IN

VIRAL LIFE CYCLE

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

STATUS INQUIRY

Sirs:

Applicants spoke with Diana of the PCT Help Desk on 14 April 2005. Diana was unable to retrieve any record of submission by international application number or attorney docket number. Enclosed, please find a copy of the submission to the USPTO filed 11 March 2005, the return express mail receipt, and the returned post card date stamped by the PCT/PTO.

Applicants have received no further correspondence or communications from the Commissioner of Patents concerning this case.

Applicant requests the current status of the application.

Respectfully submitted,

Jason C. Fedon

Agent for Applicants

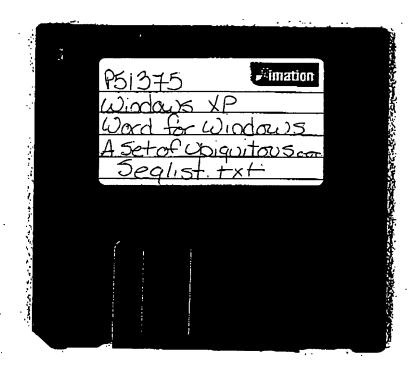
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| Amendment Response pages | ☐ Notice of Appen/Brief |
| Petition for Extension of Time | Resp. to Rest. Req pgs. |
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Attorney Docket No.: P51375

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Behrens, et al. 11 March 2005

International Serial No.: PCT/US03/28654 Group Art Unit No.: unknown

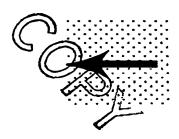
International Filing Date: 12 September 2003 Examiner: unassigned

For: A Set Of Ubiquitous Cellular Protiens Involved In Viral Life Cycle

STATEMENT TO SUPPORT FILING AND SUBMISSION IN ACCORDANCE WITH 37 CFR §§ 1.821 THROUGH 1.825

Commissioner for Patents Mail Stop: PCT P.O. Box 1450 Alexandria, Va 22313-1450

- (X) I hereby state that the contents of the paper and computer readable copies of the Sequence Listing, submitted in accordance with 37 CFR §1.821(c) and (e), respectively, are the same.
- () I hereby state that the submission filed in accordance with 37 CFR §1.821 (g) does not include new matter.
- () I hereby state that the submission filed in accordance with 37 CFR §1.821 (h) does not include new matter or go beyond the disclosure in the international application as filed.
- () I hereby state that the amendments, made in accordance with 37 CFR §1.825 (a), included in the substitute sheet(s) of the Sequence Listing are supported in the application, as filed, at pages _____. I hereby state that the substitute sheet(s) of the Sequence Listing does (do) not include new matter.
- () I hereby state that the substitute copy of the computer readable form, submitted in accordance with 37 CFR §1.825(b), is the same as the amended Sequence Listing.



Serial No.: PCT/USO. 3654 Group Art Unit No.: unknown

I hereby state that the substitute copy of the computer readable form, submitted in accordance with 37 CFR §1.825(d), is identical to that originally filed.

Respectfully submitted,

Agent for Applicants
Registration No. 48,138

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King of Prussia, PA 19406-0939
Phone (610) 270-6150
Facsimile (610) 270-5090
Sequence Listing Transmittal.doc

PATENT ATTORNEY'S DOCKET NUMBER P51375

TRANSMITTAL LETTER TO THE U.S. DESIGNATED OFFICE (DO/US) - ENTRY INTO NATIONAL STAGE UNDER 35 USC 371

INTERNATIONAL APP. NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED

PCT/US03/28654

12 September 2003

13 September 2002

TITLE OF INVENTION

A SET OF UBIQUITOUS CELLULAR PROTIENS INVOLVED IN VIRAL LIFE CYCLE

APPLICANT(S) FOR DO/US

Sven-Erik BEHRENS, Olaf ISKEN, Claus W. GRASSMANN, and Robert T. SARISKY

Commissioner for Patents

Mail Stop: PCT P.O. Box 1450

Alexandria, VA 22313-1450

ATTENTION: DO/US

CERTIFICATION UNDER 37 CFR 1.10

I hereby certify that this Transmittal Letter, Form PTO 1390 and the papers indicated as being transmitted therewith, and Post Card are being deposited with the United States Postal Service on this date March 11, 2005 in an envelope as "Express Mail Post Office to Addressee" Mailing Label Number EV608165302US addressed to the:

Commissioner for Patents, Mail Stop: PCT, P.O. Box 1450, Alexandria, VA 22313-1450.

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CUSTOMER NUMBER

| Parm PTO 1390 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE (REV 5-93) | | ATTORNEY'S DOCKET NUMBER P51375 | | |
|--|------------------------|--|--|--|
| TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED / ELECTED OFFICE (DO/EO/US) CONCERNING A FILING UNDER 35 U.S.C. 371 | | U.S. APPLICATION NO. (If known, see 37 C.F.R. 1.5) | | |
| INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PCT/US03/28654 12 September 2003 | | PRIORITY DATE CLAIMED 13 September 2002 | | |
| TITLE OF INVENTION A SET OF UBIQUITOUS CE | ELLULAR PROTIENS INVO | DLVED IN VIRAL LIFE CYCLE | | |
| APPLICANT(S) FOR DO/EO/US Sven-Erik BEHRENS, Olaf IS | SKEN, Claus W. GRASSMA | NN, and Robert T. SARISKY | | |

Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:

- 1 [x] This is a FIRST submission of items concerning a filing under 35 U.S.C. 371.
- 2. [] This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371.
- 3. [X] This express request to begin national examination procedures (35 U.S.C. 371(f)) at any time rather than delay examination until the expiration of the applicable time limit set in 35 U.S.C. 371(b) and PCT Articles 22 and 39(1).
- 4. [X] A proper Demand for International Preliminary Examination was made by the 19th month from the carliest claimed priority date.
- 5. [X] A copy of the International Application as filed (35 U.S.C. 371(c)(2))
 - a. [] is transmitted herewith (required only if not transmitted by the International Bureau).
 - b. [X] has been transmitted by the International Bureau.
 - c. [] is not required, as the application was filed in the United States Receiving Office (RO/US).
- 6. [] A translation of the International Application into English (35 U.S.C. 371(c)(2)).
- 7. [] Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))
 - a. [] are transmitted herewith (required only if not transmitted by the International Bureau).
 - b. [] have been transmitted by the International Bureau.
 - c. [] have not been made; however, the time limit for making such amendments has NOT expired.
 - d. [] have not been made and will not be made.
- 8. [] A translation of the amendments to the claims under PCT Article 19 (35 U.S. C. 371(c)(3)).
- 9. [] An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).
- A translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).

Items 11. to 16. below concern other document(s) or information included:

- 11. [] An Information Disclosure Statement under 37 C.F.R. 1.97 and 1.98; and Form PTO-1449.
- An assignment document for recording. A separate cover sheet in compliance with 37 C.F.R. 3.28 and 3.31 is included.
- 13. [X] A FIRST preliminary amendment.

(Add claim to priority via Preliminary Amendment for US originating cases only)

- 14. [] A SECOND or SUBSEQUENT preliminary amendment.
- 15. [] A substitute specification.
- 16. [] Power of attorney.
- 17. [x] An Abstract on a separate sheet of paper.
- 18. [] Copy of Form PCT/ISA/210.
- 19. [] Other items or information.

| SKB | |
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| US APPLICATION I | 10. (if known see 37 CFR 1. | 50) INTERNATIONAL PCT/US03/2 | APPLICATION NO. 8654 | ATTORNEYS DOCK P51375 | ORNEYS DOCKET NO. 1375 | |
|---|--|---|-------------------------|--------------------------|---------------------------|--|
| 20. [X] The fo | llowing fees are submit | CALCULATION | PTO USE ONLY | | | |
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| *Search Fee USPTO as | n Fee 37 CFR 1.445(a)(2) has b an International Scarchin I Search Report prepared | een paid on the internate g Authorityand and provided to the Offi | i | \$500.00 | | |
| *All other sit | uations | TOTAL OF ABOV | E CALCULATIONS = | \$1000.00 | | |
| Surcharge of \$130.00 for furnishing the oath or declaration later than \(\text{\text{\text{20}}} \) \(\text{\text{\text{30}}} \) months from the earliest claimed priority date (37 CFR 1.492(e)). | | | \$0.00 | | | |
| Claims | Number Filed | Number Extra | Rate | | | |
| Total claims | 19 - 20 = | 0 | 0 x \$50.00 | \$0.00 | | |
| Independent claims | 2- 3 = | 0 | 0x \$200.00 | \$0.00 | | |
| | ent claims (if applicabl | e) | + \$360.00 | \$0.00 | | |
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| National Stage | Application size fee - fo 50 addt'l sheets 1 x \$25 | r each additional 50 s | heets that exceed 100 | \$0.00 | | |
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| Processing fee of | of \$130.00 for furnishing this from the earliest c | laimed priority date (3 | 37 CFR 1.492(f)) + | \$ | | |
| 20 30 months from the earliest claimed priority date (37 CFR 1.492(f)) + TOTAL NATIONAL FEE = | | | \$1000.00 | | | |
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- b. Please charge my Deposit Account No. 19-2570 in the amount of \$1000.00 to cover the above fees.
- d. Seneral Authorization to charge any and all fees under 37 CFR 1.16 or 1.17, including petitions for extension of time relating to this application (37 CFR 1.136 (a)(3)).

NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the application to pending status.

SEND ALL CORRESPONDENCE TO:

GLAXOSMITHKLINE

Corporate Intellectual Property - UW2220

P.O. Box 1539

King of Prussia, PA 19406-0939

Phone (610) 270-6150

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Date of Deposit: 11 March 2005

Attorney Docket No: P51375

IN THE UNITED STATES INTERNATIONAL EXAMINING AUTHORITY

International Application No.:

PCT/US03/28654

International Filing Date:

12 September 2003

Priority Date Claimed:

13 September 2002

Applicants for DO/US:

Sven-Erik BEHRENS, Olaf ISKEN, Claus W.

GRASSMANN, and Robert T. SARISKY

Title of Invention:

A SET OF UBIQUITOUS CELLULAR PROTIENS

INVOLVED IN VIRAL LIFE CYCLE

Commissioner for Patents

Mail Stop: PCT P.O. Box 1450

Alexandria, VA 22313-1450

FIRST PRELIMINARY AMENDMENT

Sir:

Preliminary to calculating filing fees and examining this application please amend the application as follows.

Amendments to the Specification begin on page 2 of this paper.

Amendments to the Claims begin on page 3 of this paper.

Remarks/Arguments begin on page 6 of this paper.

International Applicat. No. PCT/US03/28654
International Filing Date: 12 September 2003

Amendments to the Specification

Please add the priority information paragraph to the specification by inserting the following new paragraph before the first line of the specification:

This application claims the benefit of U.S. Provisional Application No. 60/410,460, filed 13 September 2002.

An Abstract on a separate sheet is attached as required under 37 CFR 1.72(b). Please insert the attached abstract, following the claims.



International Application No. PCT/US03/28654 International Filing Date: 12 September 2003

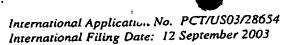
Amendments to the claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

- 1. (Original) A method for modulating viral RNA replication and translation, in a eukaryotic cell, of positive-strand viral RNA, comprising the step of contacting a viral RNA-binding protein (vRbp) with a compound that modulates an activity of said vRbp.
- 2. (Original) The method of claim 1, wherein said vRbp is selected from the group consisting of: vRbp130, vRbp120, vRbp110, vRbp84, vRbp64, and vRbp45.
- 3. (Original) The method of claim 1 wherein said activity of the vRbp is selected from the group consisting of:
 - a response to viral RNA,
 - a response to interferon induction.
 - a response to double-stranded RNA-dependent protein kinase (PKR). and
 - a response to vRbp.
- 4. (Original) The method of claim 3 wherein said response is formation of a viral:cellular ribonucleoprotein (RNP) complex.
- 5. (Original) The method of claim 4 wherein said RNP complex comprises a viral RNA:vRbp interaction.
- 6. (Original) The method of claim 5 wherein said viral RNA:vRbp interaction comprises binding of a vRbp to a viral RNA 3' untranslated region (3'UTR).
- 7. (Original) The method of claim 4 wherein said viral RNA:vRbp interaction comprises binding of a vRbp to a viral RNA 5' untranslated region (5'UTR).
- 8. (Original) The method of claim 5 wherein said 3 UTR is a UGA box consensus sequence.
- 9. (Original) The method of claim 3 wherein said response is viral circularization.

International Applicat. No. PCT/US03/28654
International Filing Date: 12 September 2003

- 10. (Original) The method of claim 9 wherein said viral circularization comprises binding of vRbp to the viral 5'UTR and 3'UTR creating a physical and functional link between both ends of the RNA.
- 11. (Original) The method of claim 9 wherein said viral circularization comprises an interaction between viral 5'UTR, 3'UTR RNA, vRbp, and cellular proteins involved in the interferon antiviral response.
- 12. (Original) The method of claim 3 wherein said response is increase in translational frameshifting that result in decreased viral replication.
- 13. (Original) The method of claim 3 wherein said response is formation of a vRbp:PKR interaction.
- 14. (Original) The method of claim 1 wherein said viral replication and translation comprises coordinated regulation of replication and translation of viral RNA.
- 15. (Original) The method of claim 1, wherein said eukaryotic cell is a mammalian cell.
- 16-17. (Cancelled)
- 18. (Original) The method of claim 1, wherein said positive strand viral RNA comprises RNA from a member of the family *Flaviviridae*.
- 19. (Original) The method of claim 1 wherein said positive strand viral RNA comprises RNA from a member of the family *Picornaviridae*.
- 20-40. (Cancelled)
- 41. (Original) A method for modulating the function of a viral 3'UTR comprising the step of contacting a 3'UTR with a compound that modulates the structure of the 3'UTR as to inhibit the interaction between 3'UTR and vRbp.
- 42. (Original) A method for screening to identify compounds that activate or that inhibit the function of vRbp which comprises a method selected from the group consisting of:



- (a) mixing a candidate compound with a solution containing a vRbp, to form a mixture, measuring activity of the vRbp in the mixture, and comparing the activity of the mixture to a standard;
- (b) detecting the effect of a candidate compound on the production of viral RNA in a eukaryotic cell, using for instance, an ELISA assay, reticulocyte lysate translation assay (luciferase RNA); and
- (c) (1) contacting a composition comprising the vRbp with the compound to be screened under conditions to permit interaction between the compound and the vRbp to assess the interaction of a compound, such interaction being associated with a second component capable of providing a detectable signal in response to the interaction of the vRbp with the compound; and
 - (2) determining whether the compound interacts with and activates or inhibits an activity of the vRbp by detecting the presence or absence of a signal generated from the interaction of the compound with the vRbp.

43-46. (Cancelled)

International Application No. PCT/US03/28654 International Filing Date: 12 September 2003



REMARKS

This Preliminary Amendment is being made upon entry of International Application No. PCT/US03/28654 into the U.S. National Phase of prosecution. In the specification, a new paragraph has been added to the first line of the specification to include the priority information. An Abstract on a separate sheet is attached as required under 37 CFR 1.72(b). Claims 16-17, 20-40, and 43-46 have been cancelled. The Applicants reserve the right to prosecute, in one or more patent applications, the claims as originally filed and/or any other claims supported by the specification. Entry of this amendment into the record is requested.

Respectfully submitted,

Jason C. Fedon
Agent for Applicants
Registration No. 48,138

GLAXOSMITHKLINE Corporate Intellectual Property UW2220 P.O. Box 1539 King of Prussia, PA 19406-0939 Phone (610) 270-6150 Facsimile (610) 270-5090

ABSTRACT OF THE DISCLOSURE

A method of modulating viral RNA replication and translation, in a eukaryotic cell, of positive-strand viral RNA, comprising the step of contacting a viral RNA-binding protein (vRbp) with a compound that modulates an activity of said protein.

SEQUENCE LISTING

<110> Behrens, Sven-Erik
 Isken, Olaf
 Grassmann, Claus W.
 Sarisky, Robert T.

<120> A Set Of Ubiquitous Cellular Proteins Involved in Viral Life Cycle

<130> P51375

<140> Unknown <141> 2005-03-11

<150> PCT/US03/28654

<151> 2003-09-12

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<213> Homo sapien

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Page 1

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1190

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SEQLIST.TXT Ser Glu Thr Gly Gly Ser His Asp Lys Arg Phe Val Met Glu Val Glu 545 550 555 560 val Asp Gly Gln Lys Phe Gln Gly Ala Gly Ser Asn Lys Lys Val Ala 565 570 575 Lys Ala Tyr Ala Ala Leu Ala Ala Leu Glu Lys Leu Phe Pro Asp Thr 580 585 590 Pro Leu Ala Leu Asp Ala Asn Lys Lys Lys Arg Ala Pro Val Pro Val 595 600 605 Arg Gly Gly Pro Lys Phe Ala Ala Lys Pro His Asn Pro Gly Phe Gly 610 620 Met GTy GTy Pro Met His Asn GTu Val Pro Pro Pro Pro Asn Leu 635 635 Gly Arg Gly Arg Gly Gly Ser Ile Arg Gly Arg Gly Arg Gly Arg Gly 645 650 655

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840
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130
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Page 7

SEQLIST.TXT Lys Ser Ile Gly Thr Ala Asn Arg Pro Met Gly Ala Gly Glu Ala Leu 260 265 270 Arg Arg Val Leu Glu Cys Leu Ala Ser Gly Ile Val Met Pro Asp Gly 275 280 285 Ser Gly Ile Tyr Asp Pro Cys Glu Lys Glu Ala Thr Asp Ala Ile Gly 290 295 300 His Leu Asp Arg Gln Gln Arg Glu Asp Ile Thr Gln Ser Ala Gln His Ala Leu Arg Leu Ala Ala Phe Gly Gln Leu His Lys Val Leu Gly Met 325 330 335 Asp Pro Leu Pro Ser Lys Met Pro Lys Lys Pro Lys Asn Glu Asn Pro 340 345 Val Asp Tyr Thr Val Gln Ile Pro Pro Ser Thr Thr Tyr Ala Ile Thr 355 Pro Met Lys Arg Pro Met Glu Glu Asp Gly Glu Glu Lys Ser Pro Ser 370 380 Lys Lys Lys Lys Ile Gln Lys Lys Glu Glu Lys Ala Glu Pro Pro 385 390 395 400 Gln Ala Met Asn Ala Leu Met Arg Leu Asn Gln Leu Lys Pro Gly Leu 415 Gln Tyr Lys Leu Val Ser Gln Thr Gly Pro Val His Ala Pro Ile Phe 420 430 Thr Met Ser Val Glu Val Asp Gly Asn Ser Phe Glu Ala Ser Gly Pro
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Met Gly Leu Bro Thr Gly Ala Gly Arg Asg Gly Arg Asg Met Gly Leu Pro Thr Gly Ala Glu Gly Arg Asp Ser Ser Lys Gly Glu 465 470 475 480 Asp Ser Ala Glu Glu Thr Glu Ala Lys Pro Ala Val Val Ala Pro Ala
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1920

1980 2040 2100

SEQLIST.TXT

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225 Thr Pro Trp Ile Leu Asp Leu Leu Gly His Tyr Ala Val Met Asn Asn 255
Pro Thr Arg Gln Pro Leu Ala Leu Asn Val Ala Tyr Arg Arg Cys Leu 265
Asp Pro Cys Glu Ser Gly Asn Phe Arg Val His Thr Val Met Thr Leu 285
Asp Pro Cys Glu Ser Gly Asn Phe Arg Val His Thr Val Met Thr Leu 290
Glu Gln Gln Asp Met Val 310
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Ser Tyr Leu Ala Ser Glu Ile Ser Thr Trp Asp Gly Val Ile Val Thr 345
Pro Ser Glu Lys Ala Tyr Glu Lys Pro Pro Glu Lys Lys Glu Gly Glu Gly Glu Gly Glu Gly Asp Ala 375
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